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09/730,380

Page 1

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L3 ANSWER 1 OF 13 CAPLUS COPYRIGHT 2001 ACS  
 AN 2000:725436 CAPLUS  
 DN 133:301171  
 TI Compositions and methods for improved delivery of ionizable hydrophobic therapeutic agents  
 IN Chen, Feng-jing; Patel, Manesh V.  
 PA Lipocine, Inc., USA  
 SO PCT Int. Appl., 99 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 FAN.CNT 1

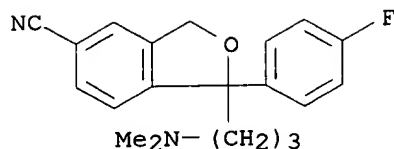
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000059475	A1	20001012	WO 2000-US7342	20000316
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 1999-287043 A 19990406				

AB The present invention is directed to a **pharmaceutical compn.** including a hydrophobic therapeutic agent having at least one ionizable functional group, and a carrier. The carrier includes an ionizing agent capable of ionizing the functional group, a surfactant, and optionally solubilizers, triglycerides, and neutralizing agents. The invention further relates to a method of prepg. such compns. by providing a compn. of an ionizable hydrophobic therapeutic agent, an ionizing agent, and a surfactant, and neutralizing a portion of the ionizing agent with a neutralizing agent. The compns. of the invention are particularly suitable for use in oral dosage forms. A carrier contg. concd.

phosphoric acid 0.025, Tween-20 0.3, Arlacel 186 0.2, sodium taurocholate 0.15, propylene glycol 0.3 g was formulated. Itraconazole was included in the carrier at 30 mg/mL for testing the stability of the itraconazole soln. upon diln. in simulated gastric fluid.

IT 59729-33-8, Citalopram  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (pharmaceutical compns. contg. hydrophobic therapeutic agents and carriers contg. ionizing agents and surfactants and triglycerides)

RN 59729-33-8 CAPLUS  
 CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



RE.CNT 3

RE

- (1) Blair; US 4306981 A 1981 CAPLUS  
 (2) Hauer; US 5342625 A 1994 CAPLUS  
 (3) Story; US 4944949 A 1990 CAPLUS

L3 ANSWER 2 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 2000:210388 CAPLUS

DN 132:246337

TI Altering patterns of activity of drug metabolizing enzymes in patients to  
 improve the effectiveness of drugs or activation of prodrugs

IN Slanetz, Alfred

PA Transgene S.A., Fr.

SO PCT Int. Appl., 23 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000017366	A2	20000330	WO 1999-EP6973	19990921
	WO 2000017366	A3	20000713		
	W: AU, CA, JP				
	RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
				US 1998-157366 A	19980921
	AU 9963280	A1	20000410	AU 1999-63280	19990921
				US 1998-157366 A	19980921
				WO 1999-EP6973 W	19990921

AB Provided are pharmaceutical compns. for the pre-treatment of a patient in need of a drug or a pro-drug wherein said drug or pro-drug is over- or under-metabolized in said patient, and uses of such **pharmaceutical compn.** The method involves either raising or lowering the level of a drug metabolizing enzyme such as a cytochrome P 450 isoenzyme assocd.

with the catabolism of a drug or the activation of a prodrug to ensure that the drug maintains a therapeutically useful level for a longer time than is typical for a given patient. The method can be extended to using a combination of enzymes that act at different stages in the treatment to maximize drug effectiveness.

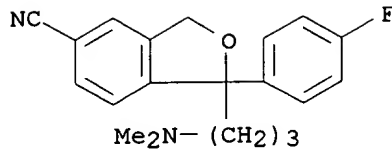
IT 59729-33-8, Citalopram

RL: BPR (Biological process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(cytochrome P 450 metab. of; altering patterns of activity of drug metabolizing enzymes in patients to improve effectiveness of drugs or activation of prodrugs)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



L3 ANSWER 3 OF 13 CAPLUS COPYRIGHT 2001 ACS  
 AN 2000:190921 CAPLUS  
 DN 132:241949  
 TI **Pharmaceutical compositions** containing NAD 299 and  
 citalopram  
 IN Evenden, John; Thorberg, Seth-Olov  
 PA Astra Aktiebolag, Swed.  
 SO PCT Int. Appl., 22 pp.  
 CODEN: PIXXD2

DT Patent  
 LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000015219	A1	20000323	WO 1999-SE1598	19990913
	W:				
	AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,				
	CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,				
	IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD,				
	MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK,				
	SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY,				
	KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,				
	ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,				
	CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
				SE 1998-3157	A 19980916
	AU 9963781	A1	20000403	AU 1999-63781	19990913
				SE 1998-3157	A 19980916
				WO 1999-SE1598	W 19990913

AB **A pharmaceutical compn.** comprising a first component  
 (a) which is (R)-3- N,N -dicyclobutylamino-8-fluoro-3,4-dihydro-2-H-1-  
 benzopyran-5-carboxamide hydrogen-(2R,3R)-tartrate monohydrate (NAD 299)  
 and a second component (b) which is citalopram, as the racemate or an  
 enantiomer thereof in the form of its free base, or a pharmaceutically  
 acceptable salt and/or solvate thereof, the prepn. thereof,

pharmaceutical

formulations contg. said compn., use of and a method of treatment of  
 affective disorders such as mood disorders and anxiety disorders with  
 said

compn. as well as a kit contg. said compn. are disclosed. S.c.  
 administration of 0.3 mg/kg NAD 299 60 min after injection of 5 mg/kg  
 citalopram to rats strongly potentiated the 5-HT elevating action of  
 citalopram vs. controls. A pharmaceutical tablet contained NAD 299 5,  
 citalopram 20, microcryst. cellulose 100, corn starch 40, povidone 4,  
 water 50, sodium starch glycolate 8, and magnesium stearate 1 mg.

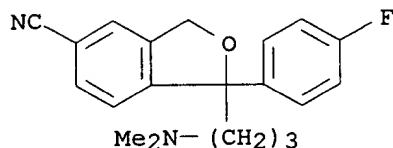
IT **59729-33-8, Citalopram**

RL: BAC (Biological activity or effector, except adverse); THU  
 (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. contg. NAD 299 and citalopram)

RN **59729-33-8 CAPLUS**

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



RE.CNT 1

RE

(1) Astra Aktiebolag; WO 9633710 A1 1996 CAPLUS

L3 ANSWER 4 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1999:282100 CAPLUS

DN 130:316651

TI Synergistic **pharmaceutical compositions** containing  
moxonidine

IN Perry, Kenneth Wayne

PA Eli Lilly and Company, USA

SO PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9920279	A1	19990429	WO 1998-US21418	19981009
	W:	AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, RO, RU, SD, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	AU 9896928	A1	19990510	US 1997-62282 P 19971017	
				AU 1998-96928	19981009
				US 1997-62282 P 19971017	
				WO 1998-US21418W	19981009
	EP 919234	A2	19990602	EP 1998-308225	19981009
	EP 919234	A3	19990825		
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
	US 6066643	A	20000523	US 1997-62282 P 19971017	
				US 1998-169369	19981009
				US 1997-62282 P 19971017	
AB	A method for producing a potentiating effect on a therapeutic action of				
an	agent which is selected from a serotonin re-uptake inhibitor, a norepinephrine re-uptake inhibitors, both a serotonin and norepinephrine re-uptake inhibitor, and an atypical antidepressant in a warm blooded mammal, comprises administering to said mammal an effective amt. of moxonidine, or a pharmaceutically acceptable salt thereof. A tablet contained moxonidine 0.300, lactose 95.700, povidone 0.700, crospovidone 3.000, magnesium stearate 0.300, hydroxypropyl Me cellulose 1.300, Et cellulose 1.200, PEG 0.250, talc 0.975, red ferric oxide 0.025, and				

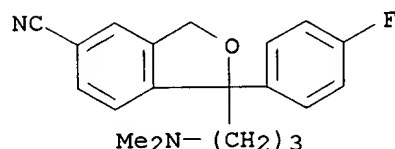
titanium dioxide 1.250 mg. Moxonidine at 0.2 mg twice daily when combined with 20 mg fluoxetine daily had synergistic effects in patients suffering major depression.

IT 59729-33-8, Citalopram

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (synergistic pharmaceutical compns. contg. moxonidine)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



RE.CNT 1

RE

(1) Armah; US 4952410 A 1990 CAPLUS

L3 ANSWER 5 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1999:81568 CAPLUS

DN 130:130004

TI **Pharmaceutical compositions** containing selective serotonin re-uptake inhibitors for the treatment and prevention of cardiac disorders using

IN Jenner, Paul Norman

PA Smithkline Beecham PLC, UK

SO PCT Int. Appl., 10 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9903469	A1	19990128	WO 1998-GB2073	19980714
	W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
				GB 1997-14841	A 19970714
	AU 9883494	A1	19990210	AU 1998-83494	19980714
				GB 1997-14841	A 19970714
				WO 1998-GB2073	W 19980714
	EP 996445	A1	20000503	EP 1998-933796	19980714
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI			
				GB 1997-14841	A 19970714
				WO 1998-GB2073	W 19980714

BR 9811004 A 20000919 BR 1998-11004 19980714  
 GB 1997-14841 A 19970714  
 WO 1998-GB2073 W 19980714  
 NO 2000000169 A 20000113 NO 2000-169 20000113  
 GB 1997-14841 A 19970714  
 WO 1998-GB2073 W 19980714

AB A method for treating and/or preventing cardiac disorders in human or non-human animals comprise administering an effective, non-toxic amt. of a serotonin re-uptake inhibitor (SSRI) or a pharmaceutically acceptable salt thereof.

A pharmaceutical tablet contained paroxetine hydrochloride hemihydrate 22.88, dibasic calcium phosphate dihydrate 244.12, hydroxypropyl methylcellulose 15.00, sodium starch glycollate 15.00, and magnesium stearate 3.00 mg. The rate of myocardial infarction for patients who were taking SSRI over 4 yr period was 0.0204 as compared to 0.0226 events/patients year exposure for the general population, showing the patients taking SSRI were statistically less likely to develop a myocardial infarction than those who did not.

IT 59729-33-8, Citalopram

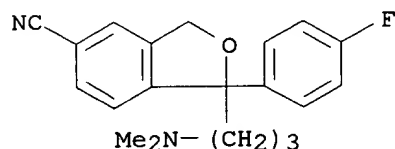
RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (pharmaceutical compns. contg. selective serotonin re-uptake

inhibitors

for treatment and prevention of cardiac disorders using)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



RE.CNT 4

RE

- (1) Carney, R; Clinical Cardiology 1997, V20(3), P196 MEDLINE
- (2) Glassman, A; J Clin Psychiatry 1993, V54(2)
- (3) Pfizer; EP 0768083 A 1997 CAPLUS
- (4) Tikal, K; Psychiatricka Lecebna Kosmonosy Ceskoslovenska Psychiatrie 1993, V89(3), P163 MEDLINE

L3 ANSWER 6 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1998:744954 CAPLUS

DN 130:17239

TI **Pharmaceutical composition** and method combining an antidepressant with an NMDA receptor antagonist, for treating neuropathic pain

IN Caruso, Frank S.

PA Algos Pharmaceutical Corp., USA

SO PCT Int. Appl., 22 pp.

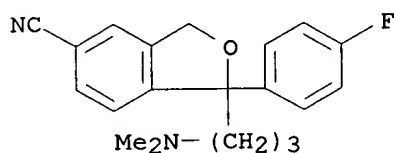
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9850044	A1	19981112	WO 1998-US9253	19980506
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9874728	A1	19981127	US 1997-45900	19970507
				AU 1998-74728	19980506
				US 1997-45900	19970507
				WO 1998-US9253	19980506
	EP 980247	A1	20000223	EP 1998-922115	19980506
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI US 1997-45900 19970507 WO 1998-US9253 19980506				
AB	The neuropathic pain alleviating effectiveness of an antidepressant is significantly potentiated by administering the antidepressant prior to, with or following the administration of a nontoxic NMDA receptor antagonist. A pharmaceutical capsule contained chlorimipramine hydrochloride 25, and dextromethorphan hydrobromide 30 mg.				
IT	<b>59729-33-8</b> , Citalopram RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical compn. and method combining antidepressant with NMDA receptor antagonist, for treating neuropathic pain)				
RN	59729-33-8 CAPLUS				
CN	5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)				



RE.CNT 8

RE

- (1) Algos Pharm Corp; WO 9627375 A 1996 CAPLUS
  - (2) Frome, B; WO 9710815 A 1997 CAPLUS
  - (3) Harris, R; WO 9640095 A 1996 CAPLUS
  - (4) Lilly Co Eli; EP 0658539 A 1995 CAPLUS
  - (5) Smith, R; WO 9609044 A 1996 CAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 7 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1998:402481 CAPLUS

DN 129:19676

TI **Pharmaceutical compositions** for the treatment of depressive disorders

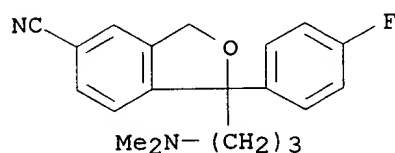
IN Medjad, Nadia; Billardon, Martine

PA UCB, S.A., Belg.  
 SO Pat. Specif. (Petty) (Aust.), 15 pp.  
 CODEN: AUXXDN

DT Patent  
 LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	AU 686084	B3	19980129	AU 1997-27539	19970626
				US 1996-672920	19960628
	US 5747494	A	19980505	US 1996-672920	19960628
AB	A method for treating a depressive disorder comprises administering to a patient in need thereof a therapeutically effective amt. of a combination (i) hydroxyzine, an individual optical isomer thereof, or a pharmaceutically acceptable salt thereof and (ii) at least one therapeutic substance which is a serotonin uptake inhibitor, an individual optical isomer thereof or a pharmaceutically acceptable salt thereof, the therapeutically effective amt. being such that the depressive disorder is treated while avoiding the nervousness, anxiety, agitation and sleep disorders assocd. with treatments using serotonin uptake inhibitors, and avoiding at the same time the loss of therapeutic effect obsd. when treatment with the classic assocn. of serotonin uptake inhibitors and benzodiazepines is used. A tablet contained fluoxetine.cntdot.HCl 10, hydroxyzine.cntdot.2HCl 25, lactose 200, and Mg stearate 1 mg. Antidepressive effects of the combination were demonstrated with rats.				
IT	59729-33-8, Citalopram RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hydroxyzine and serotonin uptake inhibitor combination for treating depressive disorder with less side effects)				
RN	59729-33-8 CAPLUS				
CN	5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)				

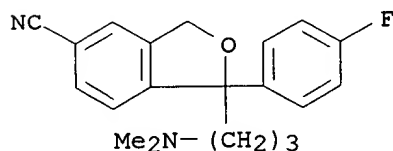


L3 ANSWER 8 OF 13 CAPLUS COPYRIGHT 2001 ACS  
 AN 1998:204419 CAPLUS  
 DN 128:261968  
 TI **Pharmaceutical composition** containing combination of atypical antipsychotic and serotonin reuptake inhibitor for treatment of psychoses  
 IN Bymaster, Franklin Porter; Perry, Kenneth Wayne; Tollefson, Gary Dennis  
 PA Eli Lilly and Co., USA  
 SO Eur. Pat. Appl., 15 pp.  
 CODEN: EPXXDW  
 DT Patent  
 LA English  
 FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	EP 830864	A1	19980325	EP 1997-307375	19970922
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
	ZA 9707967	A	19990304	US 1996-26884 P 19960923	
				ZA 1997-7967	19970904
				US 1996-26884 P 19960923	
	WO 9811897	A1	19980326	WO 1997-US15874	19970909
	W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, RO, RU, SD, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, KE, LS, MW, SD, SZ, UG, ZW, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
	AU 9744112	A1	19980414	US 1996-26884 P 19960923	
	AU 719033	B2	20000504	AU 1997-44112	19970909
				US 1996-26884 P 19960923	
				WO 1997-US15874W	19970909
	BR 9711530	A	19990824	BR 1997-11530	19970909
				US 1996-26884 P 19960923	
				WO 1997-US15874W	19970909
	CN 1230886	A	19991006	CN 1997-198113	19970909
				US 1996-26884 P 19960923	
	JP 2001503031	T2	20010306	JP 1998-514717	19970909
				US 1996-26884 P 19960923	
				WO 1997-US15874W	19970909
	US 6147072	A	20001114	US 1997-935872	19970923
				US 1996-26884 P 19960923	
	NO 9901381	A	19990322	NO 1999-1381	19990322
				US 1996-26884 P 19960923	
				WO 1997-US15874W	19970909
AB	Pharmaceutical compns. contg. combination of atypical antipsychotics and serotonin reuptake inhibitors are useful for the treatment of psychoses. Form II olanzapine (I) polymorph was prepd. by heating I at 76.degree.				
for	30 min in Et acetate and crystn. Hard gelatin capsules contained I 25, fluoxetine hydrochloride 20, starch 150, and magnesium stearate 10 mg.				
IT	59729-33-8, Citalopram				
	RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)				
	(pharmaceutical compn. contg. combination of atypical antipsychotic and serotonin reuptake inhibitor for treatment of psychoses)				
RN	59729-33-8 CAPLUS				
CN	5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)				



L3 ANSWER 9 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1998:21380 CAPLUS

DN 128:84406

TI **Pharmaceutical composition** comprising mirtazapine and one or more selective serotonin reuptake inhibitors

IN Nickolson, Victor Johannes

PA Akzo Nobel N.V., Neth.

SO Eur. Pat. Appl., 7 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 813873	A1	19971229	EP 1997-201853	19970617
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
	US 5977099	A	19991102	EP 1996-201703 A	19960619
				US 1997-876346	19970616
				EP 1996-201703 A	19960619
	CA 2208199	AA	19971219	CA 1997-2208199	19970618
				EP 1996-201703 A	19960619
	NO 9702816	A	19971222	NO 1997-2816	19970618
				EP 1996-201703 A	19960619
	BR 9703624	A	19980901	BR 1997-3624	19970618
				EP 1996-201703 A	19960619
	AU 9726129	A1	19980108	AU 1997-26129	19970619
	AU 727851	B2	20010104		
				EP 1996-201703 A	19960619
	CN 1173330	A	19980218	CN 1997-115549	19970619
				EP 1996-201703 A	19960619
	JP 10067663	A2	19980310	JP 1997-162576	19970619
				EP 1996-201703 A	19960619

AB The invention relates to a **pharmaceutical compn.** comprising mirtazapine, a selective serotonin reuptake inhibitor (SSRI) and pharmaceutically acceptable auxiliaries. In particular the SSRI is selected from fluoxetine, fluvoxamine, citalopram, cericlamine, femoxetine, sertraline, paroxetine, ifoxetine, cyanodothiepin and litoxetine. The compn. which can be used to treat depressed patients has less side effects than treatment of the patients with mirtazapine or the SSRI alone. When rats were treated with 30 mg/kg fluoxetine s.c., 80 % redn. in food consumption was reported; however, concomitant treatment with 2 mg/kg mirtazapine s.c. resulted in 40 % redn. due to partial antagonism of the appetite-inhibiting effect of fluoxetine by mirtazapine.

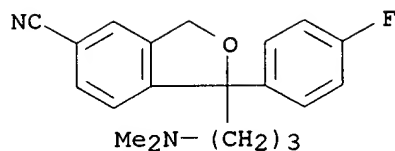
IT **59729-33-8**, Citalopram

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mirtazapine and selective serotonin reuptake inhibitor combinations for treatment of depression)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



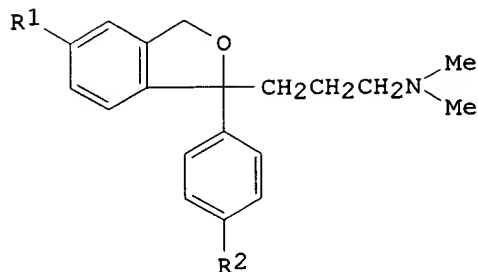
L3 ANSWER 10 OF 13 CAPLUS COPYRIGHT 2001 ACS  
 AN 1992:400917 CAPLUS  
 DN 117:917  
 TI Use of 1-(3-(dimethylamino)propyl)-1-phenylphthalans derivatives for the treatment of cerebrovascular disorders  
 IN Tanaka, Yoshiaki; Kobayashi, Naomi; Kurimoto, Tadashi; Ikeda, Yugo  
 PA Lundbeck, H., A/S, Den.  
 SO Eur. Pat. Appl., 12 pp.  
 CODEN: EPXXDW

DT Patent  
 LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 474580	A2	19920311	EP 1991-610063	19910816
EP 474580	A3	19920603		
EP 474580	B1	19940928		
R: AT, BE, CH, DE, DK, FR, GB, IT, LI, LU, NL, SE				
IL 98968	A1	19960618	DK 1990-2132	A 19900906
			IL 1991-98968	19910725
			DK 1990-2132	A 19900906
ZA 9106187	A	19920429	ZA 1991-6187	19910806
			DK 1990-2132	A 19900906
CA 2049368	AA	19920307	CA 1991-2049368	19910816
			DK 1990-2132	A 19900906
KR 9702246	B1	19970226	KR 1991-14255	19910819
			DK 1990-2132	A 19900906
AU 9182594	A1	19920312	AU 1991-82594	19910820
AU 644204	B2	19931202		
			DK 1990-2132	A 19900906
JP 04244024	A2	19920901	JP 1991-224192	19910904
JP 08005787	B4	19960124		
			DK 1990-2132	A 19900906
US 5296507	A	19940322	US 1993-1571	19930106
			DK 1990-2132	A 19900906
			US 1991-742907	B119910809

OS MARPAT 117:917  
 GI



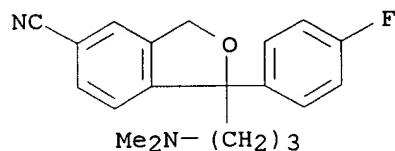
AB The title compds. [I; R1, R2 = halo, CF3, cyano, RCO (R = alkyl)] or acid addn. salts thereof are useful in the treatment of dementia, cerebrovascular disorders, and for inhibiting platelet aggregation. Citalopram (II) (40mg/kg) was i.p. injected into gerbils 30 min before carotid occlusion (5 min); 7 days later the animals were killed and surviving neurons were counted. The no. of surviving neurons was 95.8 as compared to 12.8/mm for controls. An injection soln. contained II 10, sorbitol 42.9, acetic acid 0.63, NaOH 22 mg, and water 1mL.

IT 59729-33-8, Citalopram  
RL: BIOL (Biological study)

(treatment of cerebrovascular disorders with **pharmaceutical compn.** contg.)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



L3 ANSWER 11 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1992:136283 CAPLUS

DN 116:136283

TI Pharmaceutical preparations for treatment of depression and/or migraine

IN Johnson, Edward Stewart

PA Beecham Group PLC, UK

SO PCT Int. Appl., 10 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9200103	A1	19920109	WO 1991-GB992	19910620
	W: AU, CA, JP, KR, US				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE				
				GB 1990-14354	19900628
				GB 1990-14364	19900628
				GB 1990-14365	19900628

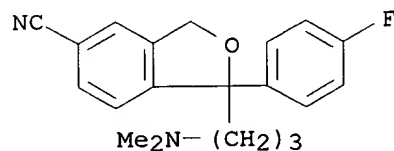
AU 9180726 A1 19920123 GB 1990-14367 19900628  
 AU 1991-80726 19910620  
 GB 1990-14354 19900628  
 GB 1990-14364 19900628  
 GB 1990-14365 19900628  
 GB 1990-14367 19900628  
 WO 1991-GB992 19910620  
 ZA 9104920 A 19920429 ZA 1991-4920 19910626  
 GB 1990-14354 19900628

AB A **pharmaceutical compn.** comprises 2-3 active ingredients selected from a 5-HT<sub>3</sub> receptor antagonist, a 5-HT reuptake inhibitor, and a 5-HT<sub>1</sub> receptor agonist, as a combined prepn. for simultaneous, sep., or sequential use in therapy.

IT **59729-33-8D**, Citalopram, mixt. with 5-HT receptor agonist and 5HT<sub>3</sub> receptor antagonists  
 RL: BIOL (Biological study)  
 (**pharmaceutical compn.** contg., for treatment of depression or migraine)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



L3 ANSWER 12 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1990:478150 CAPLUS

DN 113:78150

TI Preparation and isolation of antidepressant drug citalopram enantiomers and their **pharmaceutical compositions**

IN Boegesoe, Klaus Peter; Perregaard, Jens

PA Lundbeck, H., og Co. A/S, Den.

SO Eur. Pat. Appl., 11 pp.  
 CODEN: EPXXDW

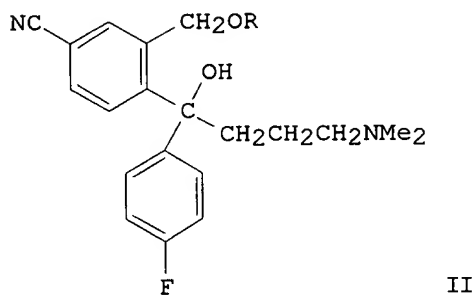
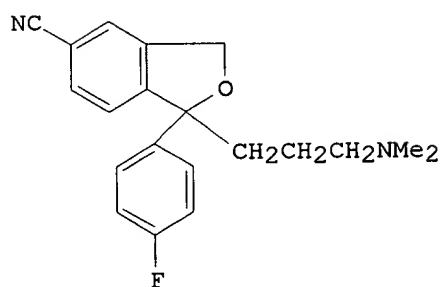
DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 347066	A1	19891220	EP 1989-305532	19890601
EP 347066	B1	19950315		
R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE				
DK 8902599	A	19891215	GB 1988-14057	A 19880614
			DK 1989-2599	19890529
			GB 1988-14057	A 19880614
IL 90465	A1	19950124	IL 1989-90465	19890530
			GB 1988-14057	A 19880614
AT 119896	E	19950415	AT 1989-305532	19890601
			GB 1988-14057	A 19880614
ES 2068891	T3	19950501	ES 1989-305532	19890601
			GB 1988-14057	A 19880614
FI 8902823	A	19891215	FI 1989-2823	19890608

FI 91527	B	19940331		
FI 91527	C	19940711		
US 4943590	A	19900724	GB 1988-14057	A 19880614
NO 8902447	A	19891215	US 1989-363589	19890608
NO 172892	B	19930614	GB 1988-14057	A 19880614
NO 172892	C	19930922	NO 1989-2447	19890613
AU 8936295	A1	19900104	GB 1988-14057	A 19880614
AU 623144	B2	19920507	AU 1989-36295	19890613
ZA 8904476	A	19900425	GB 1988-14057	A 19880614
JP 02036177	A2	19900206	ZA 1989-4476	19890613
JP 3044253	B2	20000522	GB 1988-14057	A 19880614
DK 9300115	A	19930201	JP 1989-149752	19890614
DK 170280	B1	19950724	GB 1988-14057	A 19880614
US 34712	E	19940830	DK 1993-115	19930201
FI 9401829	A	19940420	GB 1988-14057	A 19880614
CA 1339568	A1	19971202	US 1993-122009	19930914
JP 11292867	A2	19991026	GB 1988-14057	A 19880614
JP 3038204	B2	20000508	US 1989-363589	A519890608
FI 2000000507	A	20000306	FI 1994-1829	19940420
OS			GB 1988-14057	A 19880614
GI			FI 1989-2823	A 19890608
			CA 1997-617069	19970122
			GB 1988-14057	A 19880614
			CA 1989-602683	A319890613
			JP 1999-46008	19990224
			GB 1988-14057	A 19880614
			JP 1989-149752	A319890614
			FI 2000-507	20000306
			GB 1988-14057	A 19880614



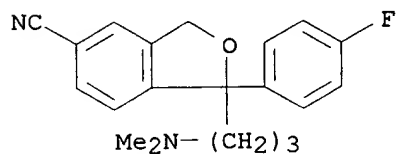
AB The title compd. (I) in pure (+)-enantiomer form and its racemic mixt., useful as antidepressants, geriatrics, or in treatment of obesity and alcoholism, are prepd. SOCl<sub>2</sub> was refluxed with a soln. of (+)-CF<sub>3</sub>CH(OMe)CO<sub>2</sub>H in CHCl<sub>3</sub> to give the acid chloride, which was dild. with CH<sub>2</sub>Cl<sub>2</sub> and treated with benzyl alc. deriv. II (R = H) and Et<sub>3</sub>N to give ester II [R = CF<sub>3</sub>CH(OMe)CO] (III) as a diastereomeric mixt., which

was purified by HPLC to give a pure enantiomer. III was dissolved in MePh and treated with Me<sub>3</sub>COK in MePh at 0.degree. to give (+)-I of 99.6% optical purity, which showed ED<sub>50</sub> of 2.0 .mu.mol/kg for 5-HTP potentiation in mice and IC<sub>50</sub> of 1.1 nM against 5-HT uptake, vs. 3.3 .mu.mol/kg and 1.8 .mu.M, resp., with (+-)-I. Similarly prepd. (-)-I showed much lower activity. Tablet, syrup, and injection formulations were given.

IT **59729-33-8P 128196-03-2P**  
 RL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. of, as antidepressant)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



RN 128196-03-2 CAPLUS

L3 ANSWER 13 OF 13 CAPLUS COPYRIGHT 2001 ACS

AN 1986:39720 CAPLUS

DN 104:39720

TI **Pharmaceutical compositions** containing unilamellar liposomes

IN Muntwyler, Rene; Hauser, Helmut

PA Ciba-Geigy A.-G., Switz.

SO Eur. Pat. Appl., 65 pp.

CODEN: EPXXDW

DT Patent

LA German

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 152379	A2	19850821	EP 1985-810050	19850211
	EP 152379	A3	19861029		
	R: AT, BE, CH, DE, FR, GB, IT, LI, LU, NL, SE				
				CH 1984-736	19840215
	ES 540372	A1	19860601	ES 1985-540372	19850213
				CH 1984-736	19840215
	CA 1246446	A1	19881213	CA 1985-474204	19850213
				CH 1984-736	19840215
	DK 8500685	A	19850816	DK 1985-685	19850214
				CH 1984-736	19840215
	AU 8538753	A1	19850822	AU 1985-38753	19850214
	AU 588798	B2	19890928		
				CH 1984-736	19840215
	ZA 8501111	A	19850925	ZA 1985-1111	19850214
				CH 1984-736	19840215
	JP 60190710	A2	19850928	JP 1985-26616	19850215

CH 1984-736 19840215

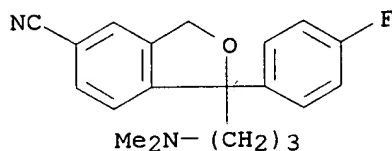
AB Aq. pharmaceutical dispersions made of unilamellar liposomes contg. an amphipathic drug and a phospholipid are given. The amphipathic drugs are quaternary ammonium compds., compds. convertible into quaternary ammonium derivs. by salt formation, .alpha.-amino acids, phosphonic acid esters, etc. Thus, 50 mg soybean lecithin was added to 20.33 mg 1-isopropylamino-3-(2-pyrrol-1-ylphenoxy)propan-2-ol-HCl [99740-06-4] in 30 mL MeOH-CHCl<sub>3</sub> (1:1) in a vial. The vial was rotated, and the film which formed was treated with 1.5 mL H<sub>2</sub>O to give a dispersion of unilamellar liposomes.

IT 59729-33-8

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(unilamellar liposome pharmaceutical compns. contg.)

RN 59729-33-8 CAPLUS

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)



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(FILE 'HOME' ENTERED AT 11:51:18 ON 23 APR 2001)

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E CITALOPRAM/CN

L1 1 S E3

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L2 782 S L1

L3 13 S PHARMACEUTICAL COMPOSITION? AND L2

=&gt; d 11

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L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2001 ACS

RN 59729-33-8 REGISTRY

CN 5-Isobenzofurancarbonitrile, 1-[3-(dimethylamino)propyl]-1-(4-fluorophenyl)-1,3-dihydro- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN (.+-.)-Citalopram

CN **Citalopram**

CN Lu 10-171

CN Nitalapram

FS 3D CONCORD

DR 128196-03-2, 103146-27-6



MF C20 H21 F N2 O

CI COM

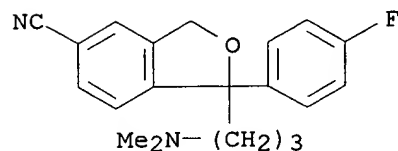
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(\*File contains numerically searchable property data)

Other Sources: EINECS\*\*, WHO

(\*\*Enter CHEMLIST File for up-to-date regulatory information)



777 REFERENCES IN FILE CA (1967 TO DATE)

5 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

780 REFERENCES IN FILE CAPLUS (1967 TO DATE)